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Response to July 16, 2003 Action

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AMENDMENTS TO THE CLAIMS: THIS LISTING OF CLAIMS REPLACES ALL PRIOR VERSIONS AND THOSE CLAIMS LISTED IN THE APPLICATION AS FILED.

Claims 1-4 (Cancelled)

5. (Withdrawn) A chemical process for preparing a cyclosporin analog of formula I as claimed in Claim 1, comprising reacting a compound of formula I, wherein A= -MeBmt-, with:
 - a. an olefin of formula $\text{CH}_2=\text{CH}-\text{X}-\text{Y}$, wherein X and Y are as defined in Claim 1, and
 - b. a catalyst;in the presence of a lithium salt in an organic solvent and optionally converting the product of said reaction into a pharmaceutically acceptable salt.
6. (Withdrawn) The process of claim 5, wherein the catalyst is Grubb's ruthenium alkylidene, Grubbs dihydroimidazole ruthenium catalyst, Schrock-Hoveyda molybdenum catalyst, Nolan's catalyst, a benzylidene catalyst or a molybdenum catalyst.
7. (Withdrawn) A chemical process for preparing a cyclosporin analog of formula I as claimed in Claim 1, comprising:
 - a. reacting a compound of formula I, wherein A= -MeBmt- with:
 - i. an olefin of formula $\text{CH}_2=\text{CH}-\text{X}-\text{Y}$, wherein X and Y are as defined in Claim 1; and
 - ii. a catalyst;in the presence of a lithium salt in an organic solvent; and
 - b. hydrogenating the product of step a in an organic solvent under hydrogen with a catalyst;and optionally converting the product of said reaction into a pharmaceutically acceptable salt.
8. (Withdrawn) The chemical process as claimed in Claim 7, wherein the catalyst in step (a) (ii) is Grubb's ruthenium alkylidene, Grubbs dihydroimidazole ruthenium catalyst,

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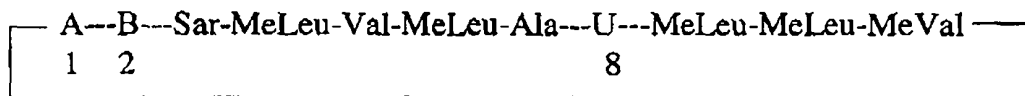
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Schrock-Hoveyda molybdenum catalyst, Nolan's catalyst, a benzyldiene catalyst or a molybdenum catalyst.

9. (Withdrawn) The chemical process as claimed in Claim 7, wherein step (b) is performed at room temperature.
10. (Withdrawn) The chemical process as claimed in Claim 9, wherein the catalyst in step (b) is Palladium on carbon or Platinum Oxide.

Claim 11 (Cancelled)

12. (Withdrawn) A method for treating autoimmune diseases in a subject, which comprises the step of administering to said subject a therapeutically effective amount of at least one cyclosporin analog of formula I as claimed in Claim 1.
13. (Withdrawn) The method of Claim 12, wherein said autoimmune disease is selected from conical cornea, keratitis, dysophia epithelialis cornea, leukoma, Mooren's ulcer, scleritis and Grave's ophthalmopathy.
14. (Withdrawn) A method for preventing organ transplantation rejection in a subject, which comprises the step of administering to said subject a therapeutically effective amount of at least one cyclosporin analog of formula I as claimed in Claim 1.
15. (Currently Amended) A cyclosporin analog of formula I or a pro-drug or a pharmaceutically acceptable salt thereof:



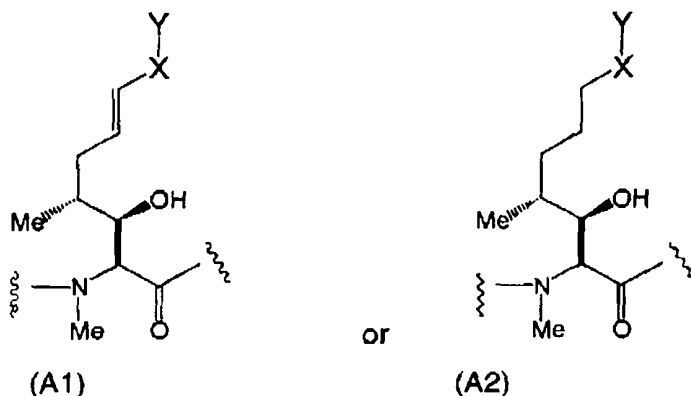
(I)

wherein

(i) A is of the formula:

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wherein:

X is absent, -C1-C6 alkyl-, or -C3-C6 cycloalkyl-;

Y is selected from the group consisting of:

- (a) aryl substituted with one or more substituents independently selected from: CN, ~~C₁-C₃-alkyl~~, C₃-C₆-alkoxy, C₁-C₆-alkoxy, C₃-C₆-alkoxy substituted with aryl, haloalkyl, thioalkoxy, amino, alkylamino, mercapto, nitro, carboxaldehyde, carboxy, alkoxycarbonyl, or carboxamide;
- (b) heteroaryl; or
- (c) substituted heteroaryl;

(ii) B is - α Abu-, -Val-, -Thr- or -Nva-; and

(iii) U is -(D)Ala-, -(D)Ser-, -[O-(2-hydroxyethyl)(D)Ser]-, -[O-(acyl)(D)Ser]- or -[O-(2-acyloxyethyl)(D)Ser]-.

16. (Currently Amended) A cyclosporin analog of claim 15 defined by formula I, wherein X is absent and Y is phenyl substituted at the ortho position with a substituent independently selected from: CN, ~~C₁-C₃-alkyl~~, C₃-C₆-alkoxy, C₁-C₆-alkoxy, C₃-C₆-alkoxy substituted with aryl, haloalkyl, thioalkoxy, amino, alkylamino, mercapto, nitro, carboxaldehyde, carboxy, alkoxycarbonyl, or carboxamide.

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17. (Currently Amended) A cyclosporin analog according to claim 15 or a pro-drug or a pharmaceutically acceptable salt thereof, selected from the group consisting of:

~~Compound of formula (I), where A=A1, X is absent and Y = (2'-Me)Ph; B is - α Abu-; and U is -(D)Ala-;~~

Compound of formula (I), where A=A1, X is absent and Y = (4'-CF₃)Ph; B is - α Abu-; and U is -(D)Ala-;

Compound of formula (I), where A=A1, X is absent and Y = (2'-OMe)Ph; B is - α Abu-; and U is -(D)Ala-;

Compound of formula (I), where A=A1, X is absent and Y = (3'-COOCH₃)Ph; B is - α Abu-; and U is -(D)Ala-;

Compound of formula (I), where A=A1, X is absent and Y = (4'-COOCH₃)Ph; B is - α Abu-; and U is -(D)Ala-;

Compound of formula (I), where A=A1, X is absent and Y = (2'-Naphthalene); B is - α Abu-; and U is -(D)Ala-;

Compound of formula (I), where A=A1, X is absent and Y = (4'-t-butyl)Ph; B is - α Abu-; and U is -(D)Ala-;

Compound of formula (I), where A=A1, X is absent and Y = (4'-AcO-)Ph; B is - α Abu-; and U is -(D)Ala-;

Compound of formula (I), where A=A1, X is absent and Y = (4'-OCH₃)Ph; B is - α Abu-; and U is -(D)Ala-;

Compound of formula (I), where A=A1, X is absent and Y = (3', 4'-OMe₂)Ph; B is - α Abu-; and U is -(D)Ala-;

~~Compound of formula (I), where A=A1, X is absent and Y = (2',5'-Me₂)Ph; B is - α Abu-; and U is -(D)Ala-;~~

Compound of formula (I), where A=A1, X is absent and Y = Pyridine; B is - α Abu-; and U is -(D)Ala-;

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Compound of formula (I), where A=A1, X is absent and Y = Pyrrole; B is $-\alpha$ Abu; and U is $-(D)Ala-$;

Compound of formula (I), where A=A1, X is absent and Y = (N-methyl) Pyrrole; B is $-\alpha$ Abu; and U is $-(D)Ala-$;

Compound of formula (I), where A=A1, X is absent and Y = Thiophene; B is $-\alpha$ Abu; and U is $-(D)Ala-$;

Compound of formula (I), where A=A1, X is absent and Y = Oxazole; B is $-\alpha$ Abu; and U is $-(D)Ala-$;

~~Compound of formula (I), where A=A2, X is absent and Y = (2'-Me)Ph; B is $-\alpha$ Abu; and U is $-(D)Ala-$;~~

Compound of formula (I), where A=A1, X is absent and Y = (S)Ph; B is $-\alpha$ Abu; and U is $-(D)Ala-$;

Compound of formula (I), where A=A1, X is absent and Y = (SO)Ph; B is $-\alpha$ Abu; and U is $-(D)Ala-$; and

Compound of formula (I), where A=A1, X is absent and Y = (SO₂)Ph; B is $-\alpha$ Abu; and U is $-(D)Ala-$.

Claim 18 (Cancelled)

19. (Previously Presented) A pharmaceutical composition, said composition comprising at least one cyclosporin analog of formula I as claimed in Claim 15, said cyclosporin analog being present alone or in combination with a pharmaceutically acceptable carrier or excipient.

20. (New) A compound according to claim 15, wherein X is absent and Y is substituted heteroaryl.

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21. (New) A compound according to claim 15, wherein X is absent and Y is (2'-methyl)furan-2-yl.